(12) NACH DEM VERTRAG ÜBER DIE INTERNATIONALE ZUSAMMENARBEIT AUF DEM GEBIET DES PATENTWESENS (PCT) VERÖFFENTLICHTE INTERNATIONALE ANMELDUNG

(19) Weltorganisation für geistiges Eigentum Internationales Büro





(43) Internationales Veröffentlichungsdatum 7. Dezember 2000 (07.12.2000)

(10) Internationale Veröffentlichungsnummer WO 00/73430 A3

(51) Internationale Patentklassifikation7: A61K 48/00. 39/395, C07K 16/42, 14/00, 16/30, A61K 39/00, A61P 35/00, 31/00, C12N 15/13, 15/10

PCT/DE00/01809

(21) Internationales Aktenzeichen: (22) Internationales Anmeldedatum:

29. Mai 2000 (29.05.2000)

(25) Einreichungssprache:

Deutsch

(26) Veröffentlichungssprache:

Deutsch

(30) Angaben zur Priorität:

199 24 405.7 27. Mai 1999 (27.05.1999)

DE 199 43 016.0 9. September 1999 (09.09.1999)

(71) Anmelder (für alle Bestimmungsstaaten mit Ausnahme von US): MAX-DELBRÜCK-CENTRUM FÜR MOLEKU-LARE MEDIZIN [DE/DE]; Robert-Rössle-Str. D-13125 Berlin (DE).

- (72) Erfinder; und
- (75) Erfinder/Anmelder (nur für US): GOLETZ, Steffen [DE/DE]; Triftstrasse 15b, D-13129 Berlin (DE).

KARSTEN, Uwe [DE/DE]; Oderbruchstrasse 29, D-10407 Berlin (DE).

- (74) Anwalt: BAUMBACH, Fritz; Robert-Rössle-Strasse 10, D-13125 Berlin (DE).
- (81) Bestimmungsstaaten (national): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW.
- (84) Bestimmungsstaaten (regional): ARIPO-Patent (GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW), eurasisches Patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), europäisches Patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI-Patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG).

Veröffentlicht:

- Mit internationalem Recherchenhericht
- (88) Veröffentlichungsdatum des internationalen Recherchenberichts: 29. März 2001

[Fortsetzung auf der nächsten Seite]

- (54) Title: VACCINES AGAINST CONFORMATION-DEPENDENT ANTIGENS AND AGAINST ANTIGENS THAT ARE NOT OR ARE NOT ONLY PROTEINS OR PEPTIDES
- (54) Bezeichnung: VAKZINE GEGEN KONFORMATIONSABHÄNGIGE ANTIGENE SOWIE GEGEN ANTIGENE, DIE KEINE ODER NICHT AUSSCHLIESSLICH PROTEINE ODER PEPTIDE SIND
- (57) Abstract: The invention relates to a method that makes it possible to use the highly effective technology of vaccination with deoxyribonucleic acid (DNA) not only on sequence epitopes of proteins or peptides, but also on conformation epitopes. The method also permits the use of DNA vaccination for antigens that are not or are only partially proteins or peptides. The preferred inventive vaccine contains a desoxyribonucleic acid (DNA) as its principal component. This desoxyribonucleic acid codes for a peptide sequence which represents the immunological imitation (mimicry) of a conformation-dependent antigen including protein conformation epitopes or of an antigen that is not or is only partially a protein or peptide. The mimicry peptide, which is also or can also be part of the inventive vaccine, is either an antiidiotypic antibody, an antibody fragment, a peptide derived therefrom or a specifically binding peptide obtained by selection from a peptide gene bank. The invention can be used in medical and veterinary medical immunology, including in the adjuvant therapy of tumor diseases.
- (57) Zusammenfassung: Die Erfindung betrifft ein Verfahren, das es erlaubt, die hocheffektive Technologie der Vakzinierung mittels Desoxyribonukleinsäure (DNA) nicht nur auf Sequenzepitope von Proteinen oder Peptiden, sondern auch auf Konformationsepitope anzuwenden. Dieses Verfahren ermöglicht darüber hinaus die Nutzung der DNA-Vakzinierung auch bei solchen Antigenen, die keine oder nur teilweise Proteine oder Peptide sind. Die bevorzugte erfindungsgemässe Vakzine enthält als wesentlichen Bestandteil eine Desoxyribonukleinsäure (DNA), die eine Peptidsequenz kodiert, welche ihrerseits die immunologische Imitation (Mimikry) eines konformationsabhängigen Antigens einschließlich Protein-Konformationsepitope oder eines Antigens, das kein oder nur teilweise Protein oder Peptid ist, darstellt. Das Mimikry-Peptid, das ebenfalls Teil der erfindungsgemäßen Vakzine ist oder sein kann, ist entweder ein antiidiotypischer Antikörper, ein Antikörperfragment, ein daraus abgeleitetes Peptid oder ein durch Selektion aus einer Peptid-Genbank erhaltenes spezifisch bindendes Peptid. Anwendungsgebiete der Erfindung sind die medizinische und die veterinärmedizinische Immunologie, darunter die adjuvante Therapie von Tumorerkrankungen.







Zur Erklärung der Zweibuchstaben-Codes, und der anderen Abkürzungen wird auf die Erklärungen ("Guidance Notes on Codes and Abbreviations") am Anfang jeder regulären Ausgabe der PCT-Gazette verwiesen.

Inte Ional Application No PCT/DE 00/01809

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 A61K48/00 A61K

A61K48/00 A61K39/00 A61K39/395 A61P35/00 C07K16/42 A61P31/00 C07K14/00 C12N15/13 C07K16/30 C12N15/10

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 7 CO7K

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

MEDLINE, LIFESCIENCES, AIDSLINE, CANCERLIT, EMBASE, CHEM ABS Data, SCISEARCH, STRAND, BIOSIS, WPI Data, EPO-Internal, PAJ

C. DOCUMEN	TS CONSIDERED	TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to daim No.
X	WO 98 00444 A (MAX DELBRUECK CT FUER MOLEKULA ;KARSTEN UWE (DE)) 8 January 1998 (1998-01-08) cited in the application the whole document	3,4,6, 10,13, 14,16
X	EP 0 508 282 A (KYOWA HAKKO KOGYO KK) 14 October 1992 (1992-10-14) column 1, line 10-15 column 2, line 45-50 example 2 claims 1,14	3,4,6, 10,13, 14,16

Y	Further documents are listed in the	continuation of box C.
^	ł	

X

Patent family members are listed in annex.

- Special categories of cited documents:
- *A* document defining the general state of the art which is not considered to be of particular relevance
- *E* earlier document but published on or after the international filing date
- "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)
- *O* document referring to an oral disclosure, use, exhibition or
- "P" document published prior to the international filing date but later than the priority date claimed
- *T* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
- *X* document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
- "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.

Date of mailing of the international search report

"&" document member of the same patent family

Date of the actual completion of the international search

.

9 November 2000

24/11/2000

Authorized officer

Name and mailing address of the ISA

Form PCT/ISA/210 (second sheet) (July 1992)

European Patent Office, P.B. 5818 Patentlaan 2 NL – 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016

Covone, M

1 84. (+31-70) 540-50

Inte ional Application No PCT/DE 00/01809

		PC1/DE 00/01809	
C.(Continu Category *	Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT egory * Citation of document, with indication, where appropriate, of the relevant passages Relevant to claim No.		
Odlegory	oranion of document, with indication, who appropriate, or the relevant passages		
X	POTTER V ET AL: "DNA vaccination with A scFv of the anti-idiotype antibody 105Ad7 induces a TH1 immune response." BRITISH JOURNAL OF CANCER, vol. 78, no. SUPPL. 2, 1998, page 18 XP000960534 Joint Meeting of the British Oncological Association, the Association of Cancer Physicians and the Royal College of Radiologists; Nottingham, England, UK; July 5-7, 1998 ISSN: 0007-0920 the whole document	1,2,5-9,	
Α	the whole document	19-26	
X	GOLLASCH H ET AL: "Identification of immunogenic peptide-mimics for the Thomsen-Friedenreich-glycoantigen." ANNALS OF HEMATOLOGY, vol. 77, no. SUPPL. 2, 1998, page S84 XP000960533 Annual Congress of the German and Austrian Societies of Hematology and Oncology; Frankfurt, Germany; October 25-28, 1998 ISSN: 0939-5555 the whole document	1-10, 13-16, 23-26	
X	PINILLA CLEMENCIA ET AL: "All-D peptides recognized by an anti-carbohydrate antibody identified from a positional scanning library." JOURNAL OF MOLECULAR BIOLOGY, vol. 283, no. 5, 13 November 1998 (1998-11-13), pages 1013-1025, XP002152467 ISSN: 0022-2836 page 283, right-hand column, line 2,3 page 1014, left-hand column, paragraph 3 page 1014, right-hand column, paragraph 3 page 1019, left-hand column, paragraph 2 page 1020, right-hand column, paragraph 3 page 1021, left-hand column, paragraph 2	3-8, 11-16	

1

Inte Ional Application No
PCT/DE 00/01809

	PCT/DE 00/01809	
	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	Relevant to claim No.
Category °	Citation of document, with indication, where appropriate, of the relevant passages	neevan to can inc.
X	LOSMAN M J ET AL: "MIMICRY OF A CARCINOEMBRYONIC ANTIGEN EPITOPE BY A RAT MONOCLONAL ANTI-IDIOTYPE ANTIBODY" INTERNATIONAL JOURNAL OF CANCER,US,NEW YORK, NY, vol. 56, no. 4, 15 February 1994 (1994-02-15), pages 580-584, XP000577759 ISSN: 0020-7136 abstract page 580, left-hand column, paragraph 2 page 580, right-hand column, paragraph 4 -page 581, left-hand column, paragraph 3	1-10,13
A	APOSTOLOPOULOS V ET AL: "Carbohydrate /peptide mimics: effect on MUC1 cancer immunotherapy." JOURNAL OF MOLECULAR MEDICINE, (1999 MAY) 77 (5) 427-36. REF: 57, XP000960532 page 429, left-hand column, paragraph 1 -right-hand column, paragraph 2 page 432, left-hand column, paragraph 3 -right-hand column, paragraph 2	1-26
A	KARSTEN UWE ET AL: "Enhanced binding of antibodies to the DTR motif of MUC1 tandem repeat peptide is mediated by site-specific glycosylation." CANCER RESEARCH, vol. 58, no. 12, 15 June 1998 (1998-06-15), pages 2541-2549, XP002112486 ISSN: 0008-5472 abstract	1-26
P,X	WO 99 40433 A (UNIV PENNSYLVANIA ; KIEBER EMMONS THOMAS (US)) 12 August 1999 (1999-08-12) page 7, line 20-29 page 8, line 24 -page 9, line 11 page 22, line 7 -page 24, line 3 page 25, line 11-16	1-18
P,X	WO 99 54457 A (POLONELLI LUCIANO ;TETI GIUSEPPE (IT); CHIRON SPA (IT)) 28 October 1999 (1999-10-28) page 2, line 12-21 page 28, line 14-21 page 31, line 12-16 page 35, line 14 -page 37, line 16	1-8, 11-16

1

Information on patent family members

Inte. :ional Application No PCT/DE 00/01809

Patent document cited in search repor	t	Publication date	Patent family member(s)	Publication date
WO 9800444	А	08-01-1998	DE 19627352 EP 0937105	
EP 0508282	Α	14-10-1992	JP 4304897 CA 2064413	77 77 77
WO 9940433	Α	12-08-1999	AU 2657599	A 23-08-1999
WO 9954457	A	28-10-1999	NONE	

Form PCT/ISA/210 (patent family annex) (July 1992)